

1 I claim:

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3 1. A surgically modified animal comprising an animal having a preoperative weight, a
4 preoperative endogenous ghrelin production and a preoperative substantially normal animal
5 gastrointestinal system that has been surgically modified, wherein said surgical modification
6 reduces the volume of the stomach of said gastrointestinal tract and reduces the digestive
7 area of said gastrointestinal tract; and, wherein postoperatively, said surgically modified animal
8 exhibits a substantially permanent reduction of said preoperative weight and a substantially
9 permanent reduction in said preoperative endogenous ghrelin production.

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11 2. The animal model of claim 1, wherein said animal is selected from the group comprising
12 murine, ovine, porcine, caprine, canine, feline, and primate animals.

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14 3. The animal model of claim 1, wherein said animal is selected from the group comprising
15 transgenic murine, ovine, porcine, caprine, canine, feline, and primate animals.

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17 4. The animal model of claim 1 wherein said animal is a Zucker rat.

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19 5. The animal model of claim of claim 1, wherein said animal is selected from the group
20 comprising genetically modified murine, ovine, porcine, caprine, canine, feline, and primate
21 animals.

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23 6. The animal model of claim 1, wherein said animal is selected from the group comprising
24 cloned murine, ovine, porcine, caprine, canine, feline, and primate animals.

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1 7. The animal model of claim 1, wherein said surgical modification is selected from the group
2 comprising bariatric surgeries, biliopancreatic diversion, gastric banding, gastric reduction,
3 gastric by-pass, gastrectomy, gastrocolostomy, gastroduodenostomy, gastroenterocolostomy,
4 gastroenteroplasty, gastroenterostomy, gastroenterotomy, gastroesophagostomy,
5 gastrogastrostomy, gastroileostomy, gastrojejunostomy, gastroplasty, vertical banded
6 gastroplasty, intestinal bypass, restriction operations, and weight-loss surgery.

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8 8. The animal model of claim 4 wherein said Zucker rat has undergone a Roux-en-Y
9 gastroplasty.

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11 9. A method for performing a Roux-en-Y gastroplasty on a Zucker Rat comprising the steps
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14 a. administering anesthesia;

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16 b. shaving and sterilizing the abdomen of said Zucker rat;

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18 c. incising the abdomen of said Zucker rat;

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20 d. identifying the terminal esophagus, lesser curvature of the stomach, and greater curvature
21 of the stomach of said Zucker rat;

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23 e. dissecting the terminal esophagus, lesser curvature of the stomach, and greater curvature of
24 the stomach of said Zucker rat free of their surrounding supportive and membranous tissues;

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1 f. creating a gastric pouch having a volume of about 20% of the presurgical stomach of said
2 Zucker rat;

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4 g. dividing the jejunum of said Zucker rat at a location about 16 cm below the ligament of
5 Treitz, thereby creating a distal portion of the divided jejunum and a proximal portion of the
6 divided jejunum of said Zucker rat;

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8 h. performing a gastrojejunostomy on said Zucker rat by anastomosing the distal portion of
9 the divided jejunum of said Zucker rat to a site on the anterior surface of the gastric fundus of
10 said Zucker rat;

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12 i. performing a jejunojejunostomy on said Zucker rat by anastomosing the proximal aspect of
13 the divided jejunum of said Zucker rat to a site at a distance of about 10 cm below the site of
14 the gastrojejunostomy;

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16 j. closing the stump of the proximal portion of the divided jejunum of said Zucker rat;

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18 k. closing the incised abdomen of said Zucker rat.

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20 10. The method of claim 9, wherein said step of creating a gastric pouch further comprises
21 the steps of:

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23 a. placing a first row of surgical staples across the stomach of said Zucker rat about 2-3 mm
24 below the gastroesophageal junction of said Zucker rat;

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1 b. placing a second row of surgical staples across the stomach of said Zucker rat about 4-5
2 mm below the gastroesophageal junction of said Zucker rat;

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4 c. reinforcing said first and second row of surgical staples with sutures.

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6 11. A method for investigating the biological mechanisms of obesity and reducing obesity
7 comprising the steps of:

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9 a. selecting a plurality of animals for confinement in a common controlled laboratory
10 environment;

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12 b. dividing said plurality of animals into at least three groups, wherein each member of a first
13 group of said plurality of animals undergoes a sham operation and is thereafter permitted to
14 consume amounts of liquid nutrients and solid nutrients ad libitum; and, wherein each
15 member of a second group of said plurality of animals undergoes a surgical modification of
16 its gastrointestinal tract and is thereafter permitted to consume amounts of liquid nutrients
17 and solid nutrients ad libitum; and, wherein each member of a third group of said plurality of
18 animals undergoes said sham operation and is thereafter permitted to feed only a mean of said
19 amounts of liquid nutrients and solid nutrients consumed by said members of said second
20 group of said plurality of animals;

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22 c. daily measuring and recording a preoperative number of calories consumed per meal, a
23 preoperative number of grams of nutrients consumed per meal, and a preoperative number of
24 meals taken by each member of each of said first, second and third groups of said plurality of
25 animals;

1 d. daily measuring and recording a preoperative body weight of each member of each of said
2 first, second and third groups of said plurality of animals;

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4 e. daily calculating and recording a preoperative total daily caloric intake and a preoperative
5 total daily number of grams of nutrients consumed by each member of each of said first,
6 second and third groups of said plurality of animals;

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8 f. performing said surgical modification of said gastrointestinal tract of each of said members
9 of said second group of said plurality of animals;

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11 g. permitting each of said members of said second group of said plurality of animals to
12 resume eating and drinking about 24 hours postoperatively;

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14 h. performing said sham operation on each of said members of said first and second groups of
15 said plurality of animals;

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17 i. permitting each of said members of said first and second groups of said plurality of animals
18 to resume eating and drinking about 24 hours postoperatively;

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20 j. feeding each member of said first and second groups of said plurality of animals a diet of
21 liquid nutrients ad libitum for about 4 postoperative days.

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23 k. additionally feeding each member of said first and second groups of said plurality of
24 animals solid nutrients ad libitum on about a 5th postoperative day and continuing until a
25 sacrifice of said members of said first and second groups.

1 1. feeding each member of said third group of said plurality of animals a diet of liquid
2 nutrients in an amount equal to a mean amount of liquid nutrients consumed by said members
3 of said second group for said about 4 postoperative days.

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5 m. additionally feeding said members of said third group of said plurality of animals an
6 amount equal to a mean amount of solid nutrients consumed by said members of said second
7 group of said plurality of animals, beginning on about a 5th postoperative day and continuing
8 until a sacrifice of said members of said third group.

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10 n. daily measuring and recording a postoperative number of calories consumed per meal, a
11 postoperative number of grams of nutrients consumed per meal, and a postoperative number
12 of meals taken by each member of each of said first, second and third groups of said plurality
13 of animals;

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15 o. daily measuring and recording a postoperative body weight of each member of each of said
16 first, second and third groups of said plurality of animals;

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18 p. daily calculating and recording a postoperative total daily caloric intake and a
19 postoperative total daily number of grams of nutrients consumed by each member of each of
20 said first, second and third groups of said plurality of animals;

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22 q. daily calculating and recording a postoperative number of calories consumed per meal, a
23 postoperative number grams of nutrients consumed per meal and a postoperative number of
24 meals taken by each member of each of said first, second and third groups of said plurality of
25 animals;

1 r. sacrificing said plurality of animals on a common postoperative day;
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3 s. postmortem, comparing said preoperative and said postoperative total daily caloric intake,
4 said preoperative and said postoperative total daily number of grams of nutrients consumed,
5 said preoperative and said postoperative number of calories consumed per meal, said
6 preoperative and said postoperative number of grams of nutrients consumed per meal, said
7 preoperative and said postoperative number of meals taken, and said preoperative and said
8 postoperative body weight for each member of said first, second and third groups of said
9 plurality of animals;

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11 t. postmortem, measuring and comparing biological factors relating to biological mechanisms
12 of obesity and reduction of obesity.

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14 12. The method of claim 11, wherein said step of selecting a plurality of animals for
15 confinement in a common controlled laboratory environment further comprises selecting a
16 plurality of animals having substantially comparable ages and preoperative body weights.

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18 13. The method of claim 11, wherein said step of selecting a plurality of animals for
19 confinement in a common controlled laboratory environment further comprises selecting said
20 plurality of animals from the group comprising murine, ovine, porcine, caprine, canine,
21 feline, and primate animals.

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23 14. The method of claim 11, wherein said step of selecting a plurality of animals for
24 confinement in a common controlled laboratory environment further comprises selecting said

1 plurality of animals from the group comprising transgenic murine, ovine, porcine, caprine,
2 canine, feline, and primate animals.

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4 15. The method of claim 11, wherein said step of selecting a plurality of animals for
5 confinement in a common controlled laboratory environment further comprises selecting said
6 plurality of animals from the group comprising genetically modified murine, ovine, porcine,
7 caprine, canine, feline, and primate animals.

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9 16. The method of claim 11, wherein said step of selecting a plurality of animals for
10 confinement in a common controlled laboratory environment further comprises selecting said
11 plurality of animals from Zucker rats.

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13 17. The method of claim 11, wherein said step of selecting a plurality of animals for
14 confinement in a common controlled laboratory environment further comprises selecting said
15 plurality of animals from the group comprising cloned murine, ovine, porcine, caprine,
16 canine, feline, and primate animals.

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18 18. The method of claim 11, wherein said step of performing said surgical modification of
19 said gastrointestinal tract of each of said members of said second group of said plurality of
20 animals further comprises selecting said surgical modification from the group comprising
21 bariatric surgeries, biliopancreatic diversion, gastric banding, gastric reduction, gastric by-
22 pass, gastrectomy, gastrocolostomy, gastroduodenostomy, gastroenterocolostomy,
23 gastroenteroplasty, gastroenterostomy, gastroenterotomy, gastroesophagostomy,
24 gastrogastrostomy, gastroileostomy, gastrojejunostomy, gastroplasty, vertical banded
25 gastroplasty, intestinal bypass, restriction operations, and weight-loss surgery.

1 19. The method of claim 11, wherein said step of dividing said plurality of animals into at
2 least three groups further comprises selecting said second group of animals from Zucker rats
3 and surgically modifying said gastrointestinal tract of each of said members of said second
4 group by means of a Roux-en-Y gastroplasty.

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6 20. The method of claim 11, wherein said sham operation comprises opening and closing the
7 abdomen of said members of said first and third groups of said plurality of animals.

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9 21. The method of claim 11, wherein said step of selecting a plurality of animals for
10 confinement in a common controlled laboratory environment further comprises confining
11 said plurality of animals to a common cage having a common standardized source of food
12 and water for a period of about one week after their selection to acclimatize them to their
13 surroundings.

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15 22. The method of claim 11, wherein said step of selecting a plurality of animals for
16 confinement in a common controlled laboratory environment further comprises providing
17 said controlled environment with an ambient temperature of about 26°C, a relative humidity
18 of about 45%, and a 12-hour light/dark cycle.

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20 23. The method of claim 11, wherein said step dividing said plurality of animals into at least
21 three groups further comprises, initially confining each member of said plurality of animals to
22 an individual cage for a period of about 1 week, equipped with a device to continuously feed,
23 measure, calculate and record said preoperative total daily number of grams of nutrients
24 consumed, said preoperative number of calories consumed per meal, said preoperative
25 number of grams of nutrients consumed per meal, said preoperative number of meals taken,

1 and said preoperative body weight for each member of said first, second and third groups of
2 said plurality of animals.

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